	CIVETIOIS COTTOCION DY THO STIC Sy Sms Branch CRE Processing Date: 21/200+
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_]. 	EN EN EN AND IN THE CONTRACT OF THE CONTRACT O
_}	
	Edited a lormat error in the Current Application Oata section, specifically:
	Edited the Current Application Data section with the actual current number. The number inputted by the applicant was [] the prior application data; or [] other
)	Added the mandatory heading and subheadings for "Current Application Data".
	Edited the 'Number of Sequences' field. The applicant spelled out a number instead of using an integer
•	Changed the spelling of a mandatory field (the headings or subheadings), specifically:
	Corrected the SEO ID NO when obviously incorrect. The sequence numbers that were edited were:
	Inserted or corrected a nucleic number at the end of a nucleic line. SEO ID NO's edited:
	Corrected subheading placement. All responses must be on the same line as each subheading. If the applicant placed a response below the subheading, this was moved to its appropriate place.
	Inserted colons after headings/subheadings. Headings edited included:+, . '
,	Deleted extra, invalid, headings-used by an applicant, specifically:
	Deleted: non-ASCII "garbage" at the beginning/end of tiles: secretary initials/filename at end of file: page numbers throughout text; other invalid text, such as
	Inserted mandatory headings, specifically:
	Corrected an obvious erro: in the response, specifically:
•	Edited identifiers where upper case is used but lower caso is required, or vice versa.
•	Corrected an orror in the Number of Sequences field, specifically:
_	1 "Hard Pago Break" code was inserted by the applicant. All occurrences had to be deleted.
	eloted ending stop codon in amino acid sequences and adjusted the *(A)Length: field accordingly (error to be a Patentin bug). Sequences corrected:
	Other: Segn 16 and 17 - moved 22237 regions up on bie
· · _	
7 7	
· <u>-</u>	

Examiner: The above corrections must be communicated to the applicant in the first Office Aciden. DO NOT send a copy of this form.

DATE: 10/10/2001

TIME: 13:44:04

Input Set : A:\PTO.AMC.txt Output Set: N:\CRF3\10102001\1671953.raw 3 <110> APPLICANT: Meares, Claude Chmura, Albert The Regents of the University of California 7 <120> TITLE OF INVENTION: Engineering Antibodies That Bind Irreversibly 9 <130> FILE REFERENCE: 023070-099120US 11 <140> CURRENT APPLICATION NUMBER: US 09/671,953 12 <141> CURRENT FILING DATE: 2000-09-27 14 <150> PRIOR APPLICATION NUMBER: US 60/156,194 15 <151> PRIOR FILING DATE: 1999-09-27 17 <150> PRIOR APPLICATION NUMBER: US 60/208,684 18 <151> PRIOR FILING DATE: 2000-05-31 20 <160> NUMBER OF SEQ ID NOS: 23 22 <170> SOFTWARE: PatentIn Ver. 2.1 24 <210> SEQ ID NO: 1 25 <211> LENGTH: 753 26 <212> TYPE: DNA 27 <213> ORGANISM: Artificial Sequence 29 <220> FEATURE: 30 <223> OTHER INFORMATION: Description of Artificial Sequence: nucleic acid that encodes Fab heavy chain of CHA255 33 <400> SEQUENCE: 1 34 agatetgaag tgaegetggt ggagtetagg ggagaeteag tgaageetgg agggtteetg 60 35 aaacteteet gtgeageete tggatteact ttaagtggtg aaaceatgte ttgggttege 120 36 cagacteegg agaagagget ggagtgggte acaaccaete ttagtggtgg tggttteace 180 37 ttctattcag ccagtgtgaa gggtcgtttc accatctcca gagacaatgc ccagaacaac 240 38 ctctatctac aactgaatag tctgaggtct gaggacacgg ccttgtattt ctgtgcaagt 300 39 categgtttg tteactgggg ceaegggaet etggteaetg tetetgeage caaaaegaeg 360 40 ggcccatcgg tetteceect ggcaccetee tecaagagea cetetggggg cacageggee 420 41 ctgggctgcc tggtcaagga ctacttcccc gaaccggtga cggtgtcgtg gaactcaggc 480 42 geoetgaeea geggegtgea eacetteeeg getgteetae agteeteaag actetaette 540 43 ctcagcagcg tggtgaccgt gcccttcaac agcttgggca cccagaccta catctgcaac 600 44 gtgaatcaca agcccagcaa caccaaggtg gacaagaaag cagagcccaa atcttgtgac 660 45 aaatctagag ggcccttcga aggtaagcct atccctaacc ctctcctcgg tctcgattct 720 753 46 acgcgtaccg gtcatcatca ccatcaccat tga 49 <210> SEQ ID NO: 2 50 <211> LENGTH: 657 51 <212> TYPE: DNA 52 <213> ORGANISM: Artificial Sequence 54 <220> FEATURE: 55 <223> OTHER INFORMATION: Description of Artificial Sequence: nucleic acid that encodes light chain mutant with Cys 56 substituted for Asn at position 97 of CHA255 59 <400> SEQUENCE: 2 60 agatetgetg ttgtgaetea ggaatetgea eteaceaeat eacetggtga aacagteaea 60 61 ctcacttqtc qctcaaqtat tggggctgtt acaactagta actatgccaa ctgggtccaa 120 62 qaaaaaccaq atcatttatt cactggtcta ataggtggta ccaataaccg ggctccgggt 180

63 qttcctqcca qattctcagg ctccctgatt ggagacaagg ctgccctcac catcacaggg 240

RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/671,953

Input Set : A:\PTO.AMC.txt

Output Set: N:\CRF3\10102001\1671953.raw

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64 gcacagactg aagatqaggc aagatatttc tgtgctctat ggtactcctg cctctgggtr 300
65 ttcqqtqqaq gaaccaaact gactgtccta agccgwackg tggctgcacc atctgtcttc 360
66 atcttcccgc catctgatga gcagttgaaa tctggaactg cctctgttgt gtgcctgctg 420
67 aataacttct atcccagaga ggccaaagta cagtggaagg tggataacgc cctccaatcg 480
68 ggtaactccc aggagagtgt cacagagcag gacagcaagg acagcaccta cagcctcagc 540
69 agcaccetga egetgageaa ageagaetae gagaaacaea aagtetaege etgegaagte 600
70 acccatcagg gcctgagyty gcccgtcaca aagagcttca acaggggaga gtgttaa
73 <210> SEQ ID NO: 3
74 <211> LENGTH: 657
75 <212> TYPE: DNA
76 <213> ORGANISM: Artificial Sequence
78 <220> FEATURE:
79 <223> OTHER INFORMATION: Description of Artificial Sequence: nucleic acid
         that encodes the unmodified light chain of CHA255
82 <400> SEQUENCE: 3
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84 ctcacttgtc gctcaagtat tggggctgtt acaactagta actatgccaa ctgggtccaa 120
85 gaaaaaccag atcatttatt cactggtcta ataggtggta ccaataaccg ggctccgggt 180
86 gttcctgcca gattctcagg ctccctgatt ggagacaagg ctgccctcac catcacaggg 240
87 qcacagactg aagatgaggc aagatatttc tgtgctctat ggtactccaa cctctgggtr 300
88 ttcggtggag gaaccaaact gactgtccta agccgwackg tggctgcacc atctgtcttc 360
89 atcttcccgc catctgatga gcagttgaaa tctggaactg cctctgttgt gtgcctgctg 420
90 aataacttct atcccagaga ggccaaagta cagtggaagg tggataacgc cctccaatcg 480
91 qqtaactccc aggagagtgt cacagagcag gacagcaagg acagcaccta cagcctcagc 540
92 agcaccctga cgctgagcaa agcagactac gagaaacaca aagtctacgc ctgcgaagtc 600
93 acccatcagg gcctgagyty gcccgtcaca aagagcttca acaggggaga gtgttaa
96 <210> SEQ ID NO: 4
97 <211> LENGTH: 657
98 <212> TYPE: DNA
99 <213> ORGANISM: Artificial Sequence
101 <220> FEATURE:
102 <223> OTHER INFORMATION: Description of Artificial Sequence: nucleic acid
          that encodes light chain mutant with Cys
103
          substituted for Ser at position 96 of CHA255
104
106 <400> SEQUENCE: 4
107 agatetgetg ttgtgaetea ggaatetgea eteaceaeat eacetggtga aacagteaea 60
108 ctcacttgtc gctcaagtat tggggctgtt acaactagta actatgccaa ctgggtccaa 120
109 gaaaaaccag atcatttatt cactggtcta ataggtggta ccaataaccg ggctccgggt 180
110 gttcctgcca gattctcagg ctccctgatt ggagacaagg ctgccctcac catcacaggg 240
111 gcacagactg aagatgaggc aagatatttc tgtgctctat ggtactgcaa cctctgggtr 300
112 ttcggtggag gaaccaaact gactgtccta agccgwackg tggctgcacc atctgtcttc 360
113 atcttcccgc catctgatga gcagttgaaa tctggaactg cctctgttgt gtgcctgctg 420
114 aataacttct atcccagaga ggccaaagta cagtggaagg tggataacgc cctccaatcg 480
115 ggtaactccc aggagagtgt cacagagcag gacagcaagg acagcaccta cagcctcagc 540
116 agcaccctga cgctgagcaa agcagactac gagaaacaca aagtctacgc ctgcgaagtc 600
117 acccatcagg gcctgagyty gcccgtcaca aagagcttca acaggggaga gtgttaa
120 <210> SEQ ID NO: 5
121 <211> LENGTH: 218
122 <212> TYPE: PRT
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Input Set : A:\PTO.AMC.txt

Output Set: N:\CRF3\10102001\I671953.raw

```
123 <213> ORGANISM: Artificial Sequence
125 <220> FEATURE:
126 <223> OTHER INFORMATION: Description of Artificial Sequence:polypeptide
          sequence of mutant light chain with Cys
          substituted for Asn at position 97 of CHA255
128
130 <220> FEATURE:
131 <221> NAME/KEY: MOD_RES
132 <222> LOCATION: (207)
133 <223> OTHER INFORMATION: Xaa = any amino acid
135 <400> SEQUENCE: 5
136 Arg Ser Ala Val Val Thr Gln Glu Ser Ala Leu Thr Thr Ser Pro Gly
                      5
139 Glu Thr Val Thr Leu Thr Cys Arg Ser Ser Ile Gly Ala Val Thr Thr
142 Ser Asn Tyr Ala Asn Trp Val Gln Glu Lys Pro Asp His Leu Phe Thr
            35
                                  40
145 Gly Leu Ile Gly Gly Thr Asn Asn Arg Ala Pro Gly Val Pro Ala Arg
                             55
                                                  60
148 Phe Ser Gly Ser Leu Ile Gly Asp Lys Ala Ala Leu Thr Ile Thr Gly
149 65
                         70
151 Ala Gln Thr Glu Asp Glu Ala Arg Tyr Phe Cys Ala Leu Trp Tyr Ser
152
154 Cys Leu Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Ser Arg
                                     105
                100
157 Thr Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln
                                120
160 Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr
                            135
                                                 140
163 Pro Arq Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser
                                             155
164 145
                        150
166 Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr
                                         170
169 Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys
                                     185
170
                180
172 His Lys Val. Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Xaa Pro
            195
173
175 Val Thr Lys Ser Phe Asn Arg Gly Glu Cys
        210
                             215
179 <210> SEQ ID NO: 6
180 <211> LENGTH: 218
181 <212> TYPE: PRT
182 <213> ORGANISM: Artificial Sequence
184 <220> FEATURE:
185 <223> OTHER INFORMATION: Description of Artificial Sequence:polypeptide
          sequence of unmodified light chain of CHA255
188 <220> FEATURE:
189 <221> NAME/KEY: MOD_RES
190 <222> LOCATION: (207)
191 <223> OTHER INFORMATION: Xaa = any amino acid
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Input Set : A:\PTO.AMC.txt

Output Set: N:\CRF3\10102001\I671953.raw

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193 <400> SEQUENCE: 6
   194 Arg Ser Ala Val Val Thr Gln Glu Ser Ala Leu Thr Thr Ser Pro Gly
                         5
                                             10
   197 Glu Thr Val Thr Leu Thr Cys Arg Ser Ser Ile Gly Ala Val Thr Thr
                    20
   198
   200 Ser Asn Tyr Ala Asn Trp Val Gln Glu Lys Pro Asp His Leu Phe Thr
                35
   203 Gly Leu Ile Gly Gly Thr Asn Asn Arg Ala Pro Gly Val Pro Ala Arg
                                55
   206 Phe Ser Gly Ser Leu Ile Gly Asp Lys Ala Ala Leu Thr Ile Thr Gly
   209 Ala Gln Thr Glu Asp Glu Ala Arg Tyr Phe Cys Ala Leu Trp Tyr Ser
                                             90
   212 Asn Leu Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Ser Arg
                   100
                                        105
   215 Thr Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln
                                   120
                                                        125
   218 Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr
                               135
           130
   221 Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser
                                                155
   224 Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr
                                                                175
                                            170
                       165
   227 Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys
                                       185
                   180
230 His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Xaa Pro
         195
                                   200
   233 Val Thr Lys Ser Phe Asn Arg Gly Glu Cys
           210
                               215
   237 <210> SEQ ID NO: 7
   238 <211> LENGTH: 218
   239 <212> TYPE: PRT
   240 <213> ORGANISM: Artificial Sequence
   242 <220> FEATURE:
   243 <223> OTHER INFORMATION: Description of Artificial Sequence:polypeptide
             sequence of mutant light chain with Cys
             substituted for Ser at position 96 of CHA255
   245
   247 <220> FEATURE:
   248 <221> NAME/KEY: MOD_RES
   249 <222> LOCATION: (207)
   250 <223> OTHER INFORMATION: Xaa = any amino acid
   252 <400> SEQUENCE: 7
   253 Arg Ser Ala Val Val Thr Gln Glu Ser Ala Leu Thr Thr Ser Pro Gly
                         5
                                             10
   256 Glu Thr Val Thr Leu Thr Cys Arg Ser Ser Ile Gly Ala Val Thr Thr
                    20
                                         25
   259 Ser Asn Tyr Ala Asn Trp Val Gln Glu Lys Pro Asp His Leu Phe Thr
   262 Gly Leu Ile Gly Gly Thr Asn Asn Arg Ala Pro Gly Val Pro Ala Arg
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Input Set : A:\PTO.AMC.txt

Output Set: N:\CRF3\10102001\1671953.raw

```
263
265 Phe Ser Gly Ser Leu Ile Gly Asp Lys Ala Ala Leu Thr Ile Thr Gly
                                              75
268 Ala Gln Thr Glu Asp Glu Ala Arg Tyr Phe Cys Ala Leu Trp Tyr Cys
271 Asn Leu Trp Val Phe Gly Gly Gly Thr Lys Leu Thr Val Leu Ser Arg
274 Thr Val Ala Ala Pro Ser Val Phe Ile Phe Pro Pro Ser Asp Glu Gln
                                120
                                                     125
275
            115
277 Leu Lys Ser Gly Thr Ala Ser Val Val Cys Leu Leu Asn Asn Phe Tyr
                            135
280 Pro Arg Glu Ala Lys Val Gln Trp Lys Val Asp Asn Ala Leu Gln Ser
                                             155
                        150
283 Gly Asn Ser Gln Glu Ser Val Thr Glu Gln Asp Ser Lys Asp Ser Thr
                                         170
                                                             175
                    165
286 Tyr Ser Leu Ser Ser Thr Leu Thr Leu Ser Lys Ala Asp Tyr Glu Lys
                180
                                     185
289 His Lys Val Tyr Ala Cys Glu Val Thr His Gln Gly Leu Ser Xaa Pro
            195
                                200
290
292 Val Thr Lys Ser Phe Asn Arg Gly Glu Cys
        210
293
296 <210> SEQ ID NO: 8
297 <211> LENGTH: 250
298 <212> TYPE: PRT
299 <213> ORGANISM: Artificial Sequence
301 <220> FEATURE:
302 <223> OTHER INFORMATION: Description of Artificial Sequence:polypeptide
          sequence of unmodified heavy chain of CHA255
305 <400> SEQUENCE: 8
306 Arg Ser Glu Val Thr Leu Val Glu Ser Arg Gly Asp Ser Val Lys Pro
309 Gly Gly Phe Leu Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Leu Ser
                                      25
                 20
312 Gly Glu Thr Met Ser Trp Val Arg Gln Thr Pro Glu Lys Arg Leu Glu
313
315 Trp Val Thr Thr Leu Ser Gly Gly Gly Phe Thr Phe Tyr Ser Ala
                             55
318 Ser Val Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Ala Gln Asn Asn
                         70
321 Leu Tyr Leu Gln Leu Asn Ser Leu Arg Ser Glu Asp Thr Ala Leu Tyr
324 Phe Cys Ala Ser His Arg Phe Val His Trp Gly His Gly Thr Leu Val
                                                         110
                100
                                     105
327 Thr Val Ser Ala Ala Lys Thr Thr Gly Pro Ser Val Phe Pro Leu Ala
330 Pro Ser Ser Lys Ser Thr Ser Gly Gly Thr Ala Ala Leu Gly Cys Leu
                             135
333 Val Lys Asp Tyr Phe Pro Glu Pro Val Thr Val Ser Trp Asn Ser Gly
                                                                 160
334 145
                        150
```

Use of n and/or Xaa has been detected in the Sequence Listing. Review the Sequence Listing to insure a corresponding explanation is presented in the <220> to <223> fields of each sequence using n or Xaa.

VERIFICATION SUMMARY

PATENT APPLICATION: US/09/671,953

DATE: 10/10/2001

TIME: 13:44:05

Input Set : A:\PTO.AMC.txt

Output Set: N:\CRF3\10102001\1671953.raw

L:172 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:5 L:230 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:6 L:289 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:7 L:600 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:20